## (FILE 'HOME' ENTERED AT 11:10:43 ON 07 JAN 2004)

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FILE 'MEDLINE, EMBASE, BIOSIS, BIOTECHDS, SCISEARCH, HCAPLUS, NTIS,
     LIFESCI' ENTERED AT 11:11:16 ON 07 JAN 2004
             11 S GLYCOSYL (A) SULFOTRANSFERASE?
L1
              8 DUP REM L1 (3 DUPLICATES REMOVED)
L2
              2 S "GST4 ALPHA"
L3
              1 DUP REM L3 (1 DUPLICATE REMOVED)
L4
L5
             53 S "GST4"
             10 S HUMAN (A) L5
L6
L7
              2 DUP REM L6 (8 DUPLICATES REMOVED)
          13789 S SULFOTRANSFERASE?
L8
L9
           5827 S HUMAN AND L8
        6311680 S CLON? OR EXPRESS? OR RECOMBINANT
L10
L11
           2771 S L9 AND L10
          13955 S "L-SELECTIN"
L12
          19920 S "P-SELECTIN".
L13
L14
          31224 S L12 OR L13
L15
            124 S L11 AND L14
             77 DUP REM L15 (47 DUPLICATES REMOVED)
L16
          59143 S "GLYCAM-1" OR "CD34" OR "MADCAM-1" OR "SGP200"
L17
             16 S L16 AND L17
L18
             16 DUP REM L18 (0 DUPLICATES REMOVED)
L19
                E ROSEN S/AU
L20
           2356 S E3
                E LEE J/AU
          13300 S E3
L21
                E HEMMERICH S/AU
            118 S E3
L22
          15770 S L21 OR L20 OR L22
L23
              3 S L5 AND L23
L24
              1 DUP REM L24 (2 DUPLICATES REMOVED)
L25
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TERMINAL (ENTER 1, 2, 3, OR ?):2

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         OCT 28
                 BIOSIS file segment of TOXCENTER reloaded and enhanced
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         DEC 08
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                 Experimental property data collected by CAS now available
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         DEC 09
                 in REGISTRY
                 STN Entry Date available for display in REGISTRY and CA/CAplus
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         DEC 09
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         DEC 17
                 DGENE: Two new display fields added
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         DEC 18
                 BIOTECHNO no longer updated
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         DEC 19
                 CROPU no longer updated; subscriber discount no longer
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                 databases
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              AND CURRENT DISCOVER FILE IS DATED 23 SEPTEMBER 2003
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=> file medline embase biosis biotechds scisearch hcaplus ntis lifesci COST IN U.S. DOLLARS SINCE FILE TOTAL FULL ESTIMATED COST

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=> dup rem 11
PROCESSING COMPLETED FOR L1
L2 8 DUP REM L1 (3 DUPLICATES REMOVED)

=> d 1-8 ibib ab

L2 ANSWER 1 OF 8 BIOTECHDS COPYRIGHT 2004 THOMSON DERWENT/ISI on STN

ACCESSION NUMBER: 2003-11055 BIOTECHDS

TITLE: New mycobacterial peptide, useful for the manufacture of a medicament for treating or preventing, or a diagnostic

reagent for identifying, mycobacterial infection;

vector plasmid-mediated recombinant protein gene transfer and expression in host cell for use in recombinant vaccine

preparation against bacterium infection

AUTHOR: JAMES B W; MARSH P; HAMPSHIRE T PATENT ASSIGNEE: MICROBIOLOGICAL RES AUTHORITY PATENT INFO: WO 2003004520 16 Jan 2003

APPLICATION INFO: WO 2002-GB3052 4 Jul 2002 PRIORITY INFO: GB 2001-23993 5 Oct 2001; GB 2001-16385 4 Jul 2001

DOCUMENT TYPE: Patent LANGUAGE: English

OTHER SOURCE: WPI: 2003-210338 [20]

AB DERWENT ABSTRACT:

NOVELTY - Isolated mycobacterial peptide (I) or its fragment, derivative or variant, encoded by a mycobacterial gene, is new.

DETAILED DESCRIPTION - (I), encoded by a mycobacterial gene (II), whose expression is induced or up-regulated under culture conditions that are nutrient-starving and that maintain mycobacterial latency. The conditions are obtainable by batch fermentation of a mycobacterium for at least 20 days post-inoculation, when compared with culture conditions

that are not nutrient-starving and that support exponential growth of the mycobacterium. INDEPENDENT CLAIMS are also included for the following: (1) identifying the mycobacterial gene; (2) an inhibitor of (I); (3) an antibody that binds to (I); (4) an attenuated mycobacterium in which a gene has been modified, which renders the mycobacterium substantially non-pathogenic; (5) an attenuated mycobacterial carrier comprising (I); (6) a DNA plasmid; (7) an RNA sequence encoded by (II); (8) an RNA vector; and (9) treating or preventing mycobacterial infection.

BIOTECHNOLOGY - Preferred Vector: The vector preferably comprises: (a) the RNA sequence encoded by (II); and (b) an integration site for a chromosome of a host cell. Preferred Inhibitor: The inhibitor is capable of preventing or inhibiting the mycobacterial peptide from exerting its native biological effect. It consists of: (a) an antibiotic capable of targeting the induced or up-regulated mycobacterial gene or its gene product; or (b) an antisense or triplex-forming nucleic acid sequence that is complementary to at least part of the inducible or up-regulatable gene. The inhibitor is capable of inhibiting a protein comprising 2-nitropropane dioxygenase, acetyltransferase, oxidoreductase, transcriptional regulator, acyl transferase, UDP-glucose dehydrogenase, phosphoribosylglycinamide formyltransferase, glutathione reductase, dihydrolipoamide, transposase, proline iminopeptidase, prolyl aminopeptidase, quinolone efflux pump, glycine betaine transporter, phosphatidylethanolamine N-methyltransferase, chalcone synthase 2, sulfotransferase, glycosyl transferase, fumarate reductase flavoprotein, aminotransferase class-II pyridoxal-phosphate, bacteriophage HK97 prohead protease, penicillin-binding protein, fatty acyl-CoA racemase, nitrilotriacetate monooxygenase, histidine kinase response regulator or hydroxymethyldihydropterine pyrophosphokinase. Preferred Gene: The gene to be modified has a wild-type coding sequence corresponding to a sequence comprising 210-4377 base pairs, fully disclosed in the specification. Preferred Carrier: The attenuated mycobacterial carrier is attenuated Salmonella, vaccine virus, fowlpox virus or Mycobacterium bovis (e.g. BCG strain). Preferred Plasmid: The DNA plasmid comprises: (a) a promoter; (b) a polyadenylation signal; and (c) a sequence that is the coding sequence of the mycobacterial gene. The promoter is cytomegalovirus and/or SV40 promoters. The polyadenylation signal consists of SV40 or bovine growth hormone polyadenylation signals. The DNA plasmid comprises 210-4377 bp. Preferred Method: Identifying the mycobacterial gene comprises: (a) culturing a first mycobacterium under culture conditions that are nutrient-starving and that maintain mycobacterial latency, where the conditions are obtainable by batch fermentation of a mycobacterium for at least 20 days post-inoculation; (b) culturing a second mycobacterium under culture conditions that are not-nutrient starving and that support exponential growth of the second mycobacterium; (c) obtaining first and second mRNA populations from the first and second mycobacteria, respectively, where the first mRNA population is obtained from the first mycobacterium and where the second mRNA is obtained from the second mycobacterium; (d) preparing first and second cDNA populations from the first and second mRNA populations, respectively, during which cDNA preparation, a detectable label is introduced into the cDNA molecules of the first and second cDNA populations; (e) isolating corresponding first and second cDNA molecules from first and second cDNA populations, respectively; (f) comparing relative amounts of label or corresponding signal emitted from the label present in the isolated first and second cDNA molecules; (g) identifying a greater amount of label or signal provided by the isolated first cDNA molecule than that provided by the isolated second cDNA molecule; and (h) identifying the first cDNA and the corresponding mycobacterial gene that is induced or up-regulated during culture of a mycobacterium under latency conditions. The corresponding first and second cDNA molecules are isolated from the first and second cDNA populations, respectively, by hybridization to an array plate containing immobilized amplified DNA sequences that have been generated from mycobacterial genomic DNA. The immobilized sequences are representative of each known gene of the

mycobacterial genome. Each representative sequence is immobilized at an identified location on the plate. The first mycobacterium is cultured under culture conditions defined by a dissolved oxygen tension of less than 10%, preferably less than 7 or 5%, air saturation when measured at 37degreesC. It is harvested at least 30, preferably 40 days post-inoculation. The culture conditions are carbon-starving to the growth of the mycobacteria. A relative induction or up-regulation is identified by a relative 3-fold, preferably 4-fold increase in the amount of label or signal provided by the isolated first cDNA molecule over that provided by the isolated second cDNA molecule. Treating or preventing mycobacterial infection comprises administering to the patient the peptide, inhibitor, antibody, attenuated mycobacterium or microbial carrier, DNA sequence or plasmid, or RNA sequence or vector.

ACTIVITY - Antibacterial. No biological data given.

MECHANISM OF ACTION - Vaccine.

USE - The peptide, inhibitor, antibody, attenuated mycobacterium or microbial carrier, DNA sequence or plasmid, or RNA sequence or vector is useful for the manufacture of a medicament for treating or preventing, or of a diagnostic reagent for identifying, mycobacterial infection (claimed).

ADMINISTRATION - The medicament is administered via intravenous, intraperitoneal or intranasal routes. No dosage given. EXAMPLE - No relevant examples given. (440 pages)

ANSWER 2 OF 8 BIOSIS COPYRIGHT 2004 BIOLOGICAL ABSTRACTS INC. on STN

DUPLICATE 1

ACCESSION NUMBER: 2002:281104 BIOSIS

DOCUMENT NUMBER:

PREV200200281104

TITLE:

Method of determining whether an agent modulates

glycosyl sulfotransferase-3.

AUTHOR(S):

Bistrup, Annette [Inventor]; Rosen, Steven D. [Inventor, Reprint author]; Tangemann, Kirsten [Inventor]; Hemmerich,

Stefan [Inventor]

CORPORATE SOURCE:

San Francisco, CA, USA

ASSIGNEE: The Regents of the University of California

PATENT INFORMATION: US 6365365 April 02, 2002

SOURCE:

Official Gazette of the United States Patent and Trademark

Office Patents, (Apr. 2, 2002) Vol. 1257, No. 1. http://www.uspto.gov/web/menu/patdata.html. e-file.

CODEN: OGUPE7. ISSN: 0098-1133.

DOCUMENT TYPE:

LANGUAGE:

Patent English

ENTRY DATE:

Entered STN: 8 May 2002

Last Updated on STN: 8 May 2002

A novel human glycosylsulfotransferase expressed in high endothelial cells (GST-3) and polypeptides related thereto, as well as nucleic acid compositions encoding the same, are provided. The subject polypeptides and nucleic acid compositions find use in a variety of applications, including research, diagnostic, and therapeutic agent screening applications. Also provided are methods of inhibiting selectin mediated binding events and methods of treating disease conditions associated therewith, particularly by administering an inhibitor of at least one of GST-3 or KSGal6ST, or homologues thereof.

ANSWER 3 OF 8 BIOTECHDS COPYRIGHT 2004 THOMSON DERWENT/ISI on STN ACCESSION NUMBER: 2001-06117 BIOTECHDS

TITLE: New glycosyl-sulfotransferases

(GST)-4-alpha, GST-4-beta and GST-6 for diagnostic and

therapeutic agent screening applications;

vector-mediated gene transfer, expression in host cell, monoclonal antibody and transgenic animal for selectin binding-inhibitor, drug screening and disease therapy,

diagnosis and gene therapy

AUTHOR:

Rosen S D; Lee J K; Hemmerich S

PATENT ASSIGNEE: Univ.California LOCATION: Oakland, CA, USA.

PATENT INFO: WO 2001006015 25 Jan 2001 APPLICATION INFO: WO 2000-US19741 19 Jul 2000

PRIORITY INFO: US 2000-593828 13 Jul 2000; US 1999-144694 20 Jul 1999

DOCUMENT TYPE: Patent LANGUAGE: English

OTHER SOURCE: WPI: 2001-138471 [14]

A glycosyl-sulfotransferase (GST) (I) selected from the group GST-4-alpha, GST-4-beta and GST-6, is claimed. Also claimed are: a fragment of (I); a DNA (II) encoding (I); a DNA or its mimetic that hybridizes to (II) or its complementary sequence; an expression cassette (III) containing a transcriptional initiation region functional in an expression host and (II) under the transcriptional regulation of the transcriptional initiation region and a transcriptional termination region; a host cell (IV) containing (III); the cellular progeny of (IV); a method of producing (I); a monoclonal antibody that specifically binds to (I); and a non-human transgenic animal model for gene function, where the animal contains an introduced alteration in a gene encoding (I). (I) is useful for inhibiting a binding event between a selectin and a

selectin ligand, which involves contacting the selectin with a non-sulfated selectin ligand. (II) encoding (I) is also useful in gene therapy to treat disorders such as acute or chronic inflammation and transplant tissue rejection and also for disease diagnosis. (44pp)

L2 ANSWER 4 OF 8 BIOSIS COPYRIGHT 2004 BIOLOGICAL ABSTRACTS INC. on STN

ACCESSION NUMBER: 2001:427531 BIOSIS DOCUMENT NUMBER: PREV200100427531

TITLE: Glycosly sulfortransferase-3.

AUTHOR(S): Bistrup, Annette [Inventor, Reprint author]; Rosen, Steven

D. [Inventor]; Hemmerich, Stefan [Inventor]

CORPORATE SOURCE: San Francisco, CA, USA

ASSIGNEE: The Regents of the University of California;

Syntex, Inc.,, Palo Alto, CA, USA

PATENT INFORMATION: US 6265192 July 24, 2001

SOURCE: Official Gazette of the United States Patent and Trademark

Office Patents, (July 24, 2001) Vol. 1248, No. 4. e-file.

CODEN: OGUPE7. ISSN: 0098-1133.

DOCUMENT TYPE: Patent LANGUAGE: English

ENTRY DATE: Entered STN: 12 Sep 2001

Last Updated on STN: 22 Feb 2002

AB A novel human glycosylsulfotransferase expressed in high endothelial cells (GST-3) and polypeptides related thereto, as well as nucleic acid compositions encoding the same, are provided. The subject polypeptides and nucleic acid compositions find use in a variety of applications, including research, diagnostic, and therapeutic agent screening applications. Also provided are methods of inhibiting selectin mediated binding events and methods of treating disease conditions associated therewith.

L2 ANSWER 5 OF 8 BIOTECHDS COPYRIGHT 2004 THOMSON DERWENT/ISI on STN

ACCESSION NUMBER: 2000-00104 BIOTECHDS

TITLE: Human and mouse glycosyl-s

Human and mouse glycosyl-sulfotransferase

-3 and related polynucleotides;

expression in mammalian host cell and antibody, used for

disease diagnosis and gene therapy

AUTHOR: Bistrup A; Rosen S D; Tangemann K; Hemmerich S

PATENT ASSIGNEE: Univ.California; Syntex

LOCATION: Oakland, CA, USA; Palo Alto, CA, USA.

PATENT INFO: WO 9949018 30 Sep 1999 APPLICATION INFO: WO 1999-US4316 26 Feb 1999

PRIORITY INFO: US 1998-190911 12 Nov 1998; US 1998-45284 20 Mar 1998

DOCUMENT TYPE: Patent

LANGUAGE:

English

OTHER SOURCE:

WPI: 1999-580442 [49]

Glycosyl-sulfotransferase-3 (GST-3, 386 or 388 amino

acids) present in other than its natural environment, is new. Also claimed are: a nucleic acid (2,032 or 1,893 bp) which encodes GST-3; an expression cassette under the control of initiation sequences and termination sequences; a host cell; a method of producing GST-3; a monoclonal antibody; a method for inhibiting the binding of a selectin and a selectin ligand; a method of inhibiting a selectin mediated binding event in a mammalian host; a method of modulating a symptom of a disease condition associated with a selectin mediated binding event; a method of diagnosing a disease state related to the abnormal levels of a sulfotransferase chosen from GST-3 and KSGal6ST; a method of determining whether an agent is capable of modulating the activity of a sulfotransferase chosen from GST-3 and KSGal6ST; and a non-human transgenic animal model for gst-3 gene function. The nucleic acid sequences, DNA probes and DNA primers derived from these, proteins and antibodies are useful in detecting homologs. The products are useful in the diagnosis of diseases associated with selectin binding interactions.

ANSWER 6 OF 8 SCISEARCH COPYRIGHT 2004 THOMSON ISI on STN

ACCESSION NUMBER:

1998:906629 SCISEARCH

THE GENUINE ARTICLE: 137GQ

TITLE:

Cloning and characterization of a human glycosyl

sulfotransferase that is restricted to high endothelial venules and confers expression of the

L-selectin recognition epitope 6-sulfo sialyl Lewis X. Bistrup A (Reprint); Bakhta S; Tangemann K; Lee J K; Gunn AUTHOR:

M D; Belov Y Y; Kannagi R; Hemmerich S; Rosen S D

CORPORATE SOURCE:

UNIV CALIF SAN FRANCISCO, SAN FRANCISCO, CA 94143; ROCHE BIOSCI, PALO ALTO, CA; AIICHI CAN RES INST, NAGOYA, AICHI,

JAPAN

COUNTRY OF AUTHOR:

USA; JAPAN

SOURCE:

MOLECULAR BIOLOGY OF THE CELL, (NOV 1998) Vol. 9, Supp.

[S], pp. 718-718.

Publisher: AMER SOC CELL BIOLOGY, PUBL OFFICE, 9650

ROCKVILLE PIKE, BETHESDA, MD 20814.

ISSN: 1059-1524.

DOCUMENT TYPE: FILE SEGMENT:

Conference; Journal LIFE

LANGUAGE:

English

REFERENCE COUNT:

ANSWER 7 OF 8 L2 ACCESSION NUMBER:

BIOSIS COPYRIGHT 2004 BIOLOGICAL ABSTRACTS INC. on STN

DOCUMENT NUMBER:

1999:17006 BIOSIS PREV199900017006

TITLE:

Cloning and characterization of a human glycosyl

sulfotransferase that is restricted to high endothelial venules and confers expression of the

L-selectin recognition epitope 6-sulfo sialyl Lewis X.

Bistrup, Annette [Reprint author]; Bakhta, Sunil;

Tangemann, Kirsten; Lee, Jin Kyu; Gunn, Michael D.; Belov, Yevgeniy Y.; Kannagi, Reiji; Hemmerich, Stefan; Rosen,

CORPORATE SOURCE:

SOURCE:

AUTHOR (S):

Univ. Calif., San Francisco, CA, USA

Molecular Biology of the Cell, (Nov., 1998) Vol. 9, No.

SUPPL., pp. 124A. print.

Meeting Info.: 38th Annual Meeting of the American Society for Cell Biology. San Francisco, California, USA. December

12-16, 1998. American Society for Cell Biology.

CODEN: MBCEEV. ISSN: 1059-1524.

DOCUMENT TYPE:

Conference; (Meeting)

Conference; Abstract; (Meeting Abstract)

LANGUAGE:

English

ENTRY DATE:

Entered STN: 20 Jan 1999

Last Updated on STN: 20 Jan 1999

L2 ANSWER 8 OF 8

SCISEARCH COPYRIGHT 2004 THOMSON ISI on STN

ACCESSION NUMBER:

1998:810754 SCISEARCH

THE GENUINE ARTICLE: 130CC TITLE: Cloni:

Cloning and functional characterization of a human

glycosyl sulfotransferase, that is

highly restricted to high endothelial venules and confers expression of the L-selectin recognition epitope 6-sulfo

sialyl Lewis x.

**AUTHOR:** 

Hemmerich S (Reprint); Bistrup A; Bakhta S; Gunn M D;

Kannagi R; Rosen S D

CORPORATE SOURCE:

ROCHE BIOSCI, PALO ALTO, CA; UNIV CALIF SAN FRANCISCO, SAN

FRANCISCO, CA 94143; AIICHI CANC RES INST, NAGOYA, AICHI,

JAPAN

COUNTRY OF AUTHOR:

USA; JAPAN

SOURCE:

GLYCOBIOLOGY, (NOV 1998) Vol. 8, No. 11, pp. 29-29.

Publisher: OXFORD UNIV PRESS, GREAT CLARENDON ST, OXFORD

OX2 6DP, ENGLAND. ISSN: 0959-6658.

DOCUMENT TYPE:

Conference; Journal

FILE SEGMENT: LANGUAGE:

English

REFERENCE COUNT:

0

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FILE 'MEDLINE, EMBASE, BIOSIS, BIOTECHDS, SCISEARCH, HCAPLUS, NTIS, LIFESCI' ENTERED AT 11:11:16 ON 07 JAN 2004

L1 11 S GLYCOSYL (A) SULFOTRANSFERASE?

L2 8 DUP REM L1 (3 DUPLICATES REMOVED)

=> s "gst4 alpha"

L3 2 "GST4 ALPHA"

=> dup rem 13

PROCESSING COMPLETED FOR L3

L4 1 DUP REM L3 (1 DUPLICATE REMOVED)

=> d all

L4 ANSWER 1 OF 1 EMBASE COPYRIGHT 2004 ELSEVIER INC. ALL RIGHTS RESERVED.

ON STN DUPLICATE 1

AN 2002402885 EMBASE

TI Enzymatic synthesis in vitro of the disulfated disaccharide unit of corneal keratan sulfate.

AU Akama T.O.; Misra A.K.; Hindsgaul O.; Fukuda M.N.

CS T.O. Akama, Glycobiology Program, Burnham Institute, 10901 N. Torrey Pines Rd., San Diego, CA 92037, United States. takama@burnham-inst.org

SO Journal of Biological Chemistry, (8 Nov 2002) 277/45 (42505-42513). Refs: 54

ISSN: 0021-9258 CODEN: JBCHA3

CY United States

DT Journal; Article

FS 029 Clinical Biochemistry

LA English

SL English

AB Among the enzymes of the carbohydrate sulfotransferase family, human corneal GlcNAc 6-O-sulfotransferase (hCGn6ST, also known as human

GlcNAc6ST-5/GST4.beta.) and human intestinal GlcNAc 6-0-sulfotransferase (hIGn6ST or human GlcNAc6ST-3/GST4.alpha.) are highly homologous. In the mouse, intestinal GlcNAc 6-O-sulfotransferase (mIGn6ST or mouse GlcNAc6ST-3/GST4) is the only orthologue of hCGn6ST and hIGn6ST. In the previous study, we found that hCGn6ST and mIGn6ST, but not hIGn6ST, have sulfotransferase activity to produce keratan sulfate (Akama, T. O., Nakayama, J., Nishida, K., Hiraoka, N., Suzuki, M., McAuliffe, J., Hindsgaul, O., Fukuda, M., and Fukuda, M. N. (2001) J. Biol. Chem. 276, 16271-16278). In this study, we analyzed the substrate specificities of these sulfotransferases in vitro using synthetic carbohydrate substrates. We found that all three sulfotransferases can transfer sulfate to the nonreducing terminal GlcNAc of short carbohydrate substrates. Both hCGn6ST and mIGn6ST, but not hIGn6ST, transfer sulfate to longer carbohydrate substrates that have poly-N-acetyllactosamine structures, suggesting the involvement of hCGn6ST and mIGn6ST in production of keratan sulfate. To clarify further the involvement of hCGn6ST in biosynthesis of keratan sulfate, we reconstituted the biosynthetic pathway in vitro by sequential enzymatic treatment of a synthetic carbohydrate substrate. Using four enzymes, .beta.1,4-galactosyltransferase-I, .beta.1,3-N-acetylglucosaminyltransferase-2, hCGn6ST, and keratan sulfate Gal 6-0-sulfotransferase, we were able to synthesize in vitro a product that conformed to the basic structural unit of keratan sulfate. Based on these results, we propose a biosynthetic pathway for N-linked keratan sulfate on corneal proteoglycans.

Medical Descriptors: \*protein synthesis \*enzyme mechanism protein family cornea sequence homology amino acid sequence enzyme activity enzyme substrate enzyme specificity in vitro study biosynthesis protein structure human human cell article priority journal Drug Descriptors: \*disaccharide \*keratan sulfate sulfotransferase sulfate poly n acetylactosamine beta 1,4 galactosyltransferase 1 beta 1,3 n acetylglucosaminyltransferase 2 keratan sulfate galactosyl 6 o sulfotransferase transferase proteoglycan unclassified drug (keratan sulfate) 69992-87-6, 9056-36-4; (sulfotransferase) 9023-09-0; (sulfate) 14808~79-8; (transferase) 9047-61-4

=> d his

(FILE 'HOME' ENTERED AT 11:10:43 ON 07 JAN 2004)

FILE 'MEDLINE, EMBASE, BIOSIS, BIOTECHDS, SCISEARCH, HCAPLUS, NTIS, LIFESCI' ENTERED AT 11:11:16 ON 07 JAN 2004

11 S GLYCOSYL (A) SULFOTRANSFERASE?

8 DUP REM L1 (3 DUPLICATES REMOVED)  $L_2$ 2 S "GST4 ALPHA"  $L_3$ 1 DUP REM L3 (1 DUPLICATE REMOVED) T.4 => s "GST4" 53 "GST4" 1.5 => s human (a) 15 10 HUMAN (A) L5 L6 => dup rem 16 PROCESSING COMPLETED FOR L6 2 DUP REM L6 (8 DUPLICATES REMOVED) => d 1-2 ibib abDUPLICATE 1 ANSWER 1 OF 2 MEDLINE on STN ACCESSION NUMBER: 2001205848 MEDLINE DOCUMENT NUMBER: 21096027 PubMed ID: 11181564 Chromosomal localization and genomic organization for the TITLE: galactose/ N-acetylgalactosamine/N-acetylglucosamine 6-0-sulfotransferase gene family. Hemmerich S; Lee J K; Bhakta S; Bistrup A; Ruddle N R; **AUTHOR:** Rosen S D Department of Respiratory Diseases, Roche Bioscience, Palo CORPORATE SOURCE: Alto, CA 94304, USA. CONTRACT NUMBER: RO1GM5741 (NIGMS) GLYCOBIOLOGY, (2001 Jan) 11 (1) 75-87. SOURCE: Journal code: 9104124. ISSN: 0959-6658. PUB. COUNTRY: England: United Kingdom Journal; Article; (JOURNAL ARTICLE) DOCUMENT TYPE: English LANGUAGE: FILE SEGMENT: Priority Journals GENBANK-AF176838; GENBANK-AF280086; GENBANK-AF280087; OTHER SOURCE: GENBANK-AF280088; GENBANK-AF280089; GENBANK-AI824100 ENTRY MONTH: 200106 ENTRY DATE: Entered STN: 20010611 Last Updated on STN: 20010611 Entered Medline: 20010607 AB The galactose/N-acetylgalactosamine/N-acetylglucosamine 6-O-sulfotransferases (GSTs) are a family of Golgi-resident enzymes that transfer sulfate from 3'phosphoadenosine 5'phospho-sulfate to the 6-hydroxyl group of galactose, N-acetylgalactosamine, or N-acetylglucosamine in nascent glycoproteins. These sulfation modifications are functionally important in settings as diverse as cartilage structure and lymphocyte homing. To date six members of this gene family have been described in human and in mouse. We have determined the chromosomal localization of these genes as well as their genomic organization. While the broadly expressed enzymes implicated in proteoglycan biosynthesis are located on different chromosomes, the highly tissue specific enzymes GST-3 and 4 are encoded by genes located both in band q23.1--23.2 on chromosome 16. In the mouse, both genes reside in the syntenic region 8E1 on chromosome 8. This cross-species conserved clustering is suggestive of related functional roles for these genes. human GST4 locus actually contains two highly similar open reading frames (ORF) that are 50 kb apart and encode two highly similar enzyme isoforms termed GST-4 alpha and GST-4 beta. All genes except GST0 (chondroitin 6-O-sulfotransferase) contain intron-less ORFs. With one exception these are fused directly to sequences encoding the 3' untranslated regions (UTR) of the respective mature mRNAs. The 5' UTRs of these mRNAs are usually encoded by a number of short exons 5' of the

respective ORF. 5'UTRs of the same enzyme expressed in different cell types are sometimes derived from different exons located upstream of the ORF. The genomic organization of the GSTs resembles that of certain

glycosyltransferase gene families.

L7 ANSWER 2 OF 2 MEDLINE on STN DUPLICATE 2

ACCESSION NUMBER: 95251394 MEDLINE

DOCUMENT NUMBER: 95251394 PubMed ID: 7733673

TITLE: Cloning and expression of a cDNA for mu-class glutathione

S-transferase from rabbit liver.

AUTHOR: Lee S H; Lee S H; Han J S; Kim Y S; Koh J K

CORPORATE SOURCE: Department of Biochemistry, College of Medicine, Hanyang

University, Seoul, Korea.

SOURCE: ARCHIVES OF BIOCHEMISTRY AND BIOPHYSICS, (1995 Apr 20) 318

(2) 424-9.

Journal code: 0372430. ISSN: 0003-9861.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals OTHER SOURCE: GENBANK-L23766

ENTRY MONTH: 199505

ENTRY DATE: Entered STN: 19950608

Last Updated on STN: 19980206 Entered Medline: 19950526

A mu-class glutathione S-transferase (GST) cDNA clone, pHMB1, from rabbit AΒ liver has been constructed, using a 748-base-pair fragment of GST Yb1 cDNA as a probe. The nucleotide sequence of pHMB1 has been determined, and the complete amino acid sequence has been deduced. Recombinant clone pHMB1 contains a cDNA insert of 1443 base pairs with 654 nucleotides of open reading frame, 33 nucleotides of 5'-untranslated region, and 756  $\,$ nucleotides of 3'-untranslated region. The open reading frame encodes a polypeptide (rbGST mu I) comprising 218 amino acids with molecular weight of 25,417. Compared to published mu-class GST sequences, rbGST mu I is 73 and 77% identical to rat Yb1 and human GST4 in amino acid sequence, respectively. The pHMB1 was expressed in Escherichia coli using expression vector pIH821 and the expressed GST was purified as a single band on polyacrylamide gel electrophoresis by maltose- and glutathione-affinity column chromatography. Rabbit liver GST protein expressed by this system was catalytically active. The functional characterization was done on the expressed protein. The rabbit liver GST expressed in E. coli showed greater activity toward 1,2-dichloro-4nitrobenzene than mu-class isozymes in rabbit hepatic tissue (T. Primiano and R.F. Novak (1993) Arch. Biochem. Biophys. 301, 404-410). Enzymatic activity of expressed protein toward the substrate 1-chloro-2,4dinitrobenzene was inhibited by triethyltin bromide, Cibacron blue, triphenyltin chloride, bromosulfophthalein, and hematin. RNA blot hybridization demonstrated that the pHMB1 mRNA was well expressed in rabbit liver, brain, and kidney.

## => d his

(FILE 'HOME' ENTERED AT 11:10:43 ON 07 JAN 2004)

FILE 'MEDLINE, EMBASE, BIOSIS, BIOTECHDS, SCISEARCH, HCAPLUS, NTIS, LIFESCI' ENTERED AT 11:11:16 ON 07 JAN 2004

L1 11 S GLYCOSYL (A) SULFOTRANSFERASE?

L2 8 DUP REM L1 (3 DUPLICATES REMOVED)

L3 2 S "GST4 ALPHA"

L4 1 DUP REM L3 (1 DUPLICATE REMOVED)

L5 53 S "GST4"

L6 10 S HUMAN (A) L5

L7 2 DUP REM L6 (8 DUPLICATES REMOVED)

<sup>=&</sup>gt; s sulfotransferase?

=> s human and 18

L9 5827 HUMAN AND L8

=> s clon? or express? or recombinant

4 FILES SEARCHED...

L10 6311680 CLON? OR EXPRESS? OR RECOMBINANT

=> s 19 and 110

L11 2771 L9 AND L10

=> s "l-selectin"

L12 13955 "L-SELECTIN"

=> s "p-selectin"

L13 19920 "P-SELECTIN"

=> s 112 or 113

L14 31224 L12 OR L13

=> s l11 and l14

L15 124 L11 AND L14

=> dup rem 115

PROCESSING COMPLETED FOR L15

L16 77 DUP REM L15 (47 DUPLICATES REMOVED)

=> s "GLYCAM-1" or "CD34" or "MAdCAM-1" or "Sgp200"

L17 59143 "GLYCAM-1" OR "CD34" OR "MADCAM-1" OR "SGP200"

=> s 116 and 117

L18 16 L16 AND L17

=> dup rem 118

PROCESSING COMPLETED FOR L18

L19 16 DUP REM L18 (0 DUPLICATES REMOVED)

=> d 1-16 ibib ab

L19 ANSWER 1 OF 16 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER:

2002:251882 HCAPLUS

DOCUMENT NUMBER:

136:291000

TITLE:

Screening of novel human glycosyl sulfotransferase expressed in high

endothelial cells (HEC) (GST-3, HEC-GlcNAc6ST)

inhibitors

INVENTOR(S):

Bistrup, Annette; Rosen, Steven D.; Tangemann,

Kirsten; Hemmerich, Stefan

PATENT ASSIGNEE(S):

The Regents of the University of California, USA

U.S., 38 pp., Cont.-in-part of U.S. Ser. No. 45,284.

CODEN: USXXAM

DOCUMENT TYPE:

Patent English

LANGUAGE:

SOURCE:

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

PATENT NO.	KIND	DATE				CATI			DATE			
US 6365365	B1	20020402		US	19:	98-19	9091:	l	1998.	1112		
US 6265192	B1	20010724		US	19:	98-4	5284		19980	0320		
CA 2322779	AA	19990930		CF	19	99-23	3227	79	19990	0226		
WO 9949018	A1	19990930		WC	19:	99-U	34316	5	1999	0226		
W: AL, AM,	AT, AU	, AZ, BA,	BB,	BG,	BR,	BY,	CA,	CH,	CN,	CU,	CZ,	DE,
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KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN,
             MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM,
             TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU,
             TJ, TM
         RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK,
             ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG,
             CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                                           AU 1999-27945
                                                            19990226
     AU 9927945
                       Α1
                            19991018
                            20030904
     AU 764852
                       B2
                                           EP 1999-908538
                            20001227
                                                            19990226
     EP 1062326
                       Α1
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, FI
                                           JP 2000-537979
                                                            19990226
                            20020312
     JP 2002507409
                       T2
                                           US 2001-816825
                                                            20010322
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                       A1
                            20011213
                                           US 2001-7262
                                                            20011108
     US 2002164748
                       A1
                            20021107
                                        US 1998-45284
                                                         A2 19980320
PRIORITY APPLN. INFO.:
                                        US 1998-190911
                                                         Α
                                                            19981112
                                                         W
                                                            19990226
                                        WO 1999-US4316
     Use of a novel human glycosyl sulfotransferase
AB
     expressed in high endothelial cells (HEC) (GST-3 or HEC-GlcNAc6ST)
     for screening inhibitors as therapeutic agent is provided. Full-length
     cDNAs contg. the two contigs and predicting CS6T/KSST homologs were
     obtained by screening a human fetal brain .lambda.ZAP cDNA
     library (Stratagene, La Jolla, Calif.) with labeled 700-800 bp restriction
     fragments (from EST 2 for contig 1 and from EST 5 for contig 2). The
     proteins encoded by these cDNAs were designated as GST 1 and GST 2, where
     GST denotes "glycosylsulfotransferase." GST 1 has been independently
     cloned and assigned the name "KSGal6ST by Fukuta et al., J. Biol.
     Chem. (1997) 272: 32321-8. ESTs potentially coding for novel
     human glycosyl sulfotransferases other than GST-1&2 were
     identified through a secondary homol. screen, in which the peptide
     sequences of GST-1 and GST-2 were used as template in two parallel TBLASTN
     searches against a public (dbest) and a private genomic database (Lifeseq,
     Incyte Pharmaceuticals, Palo Alto, Calif.). Three cDNA clones
     which encode three different human homologs for C6ST/KSST have
     been obtained. The predicted GST proteins are type 2 membrane proteins
     411, 484, and 386 amino acids in length, resp. Each has a relatively
     short transmembrane domain and a short amino terminal cytoplasmic tail.
     GST-1 is the same as the sulfotransferase reported by Fukuta et
     al. supra (1997) and named KSGal6ST. GST-3 (HEC-GlcNAc6ST), is a novel
     GlcNAc-6-sulfotransferase. The novel human
     glycosylsulfotransferase enzyme of the subject invention has been named
     human glycosyl sulfotransferase 3 or huGST-3 or
     HEC-GlcNAc6ST. HuGST-3 is capable of sulfating selectin ligands,
     particularly L-selectin ligands, e.g., GlyCAM
     -1. Donor compds. from which huGST-3 obtains sulfate groups for
     transfer to acceptor ligand compds. include 3'-phosphoadenosine
     5'-phosphosulfate (PAPS) and the like. Selectin ligands capable of being
     sulfated through huGST-3 action include E-, P- and L-
     selectin ligands, particularly L-selectin
     ligands, such as GlyCAM-1, CD34,
     MAdCAM-1, Sgp200, podocalyxin, and the like.
     huGST-3 is strongly predicted to have GlcNAc6-O-sulfotransferase
     (N-actylglucosamine-6-0-sulfotransferase) activity.
     Human GST-3 is a 386 amino acid protein having an amino acid
     sequence as shown in FIG. 1 and identified as SEQ ID NO:01.
                               THERE ARE 37 CITED REFERENCES AVAILABLE FOR THIS
REFERENCE COUNT:
                         37
                               RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
                        MEDLINE on STN
     ANSWER 2 OF 16
                                   MEDLINE
ACCESSION NUMBER:
                    2002463169
```

DOCUMENT NUMBER: 22194291 PubMed ID: 12068018
TITLE: Distinct sulfation requirements of selectins disclosed using cells that support rolling mediated by all three

selectins under shear flow. L-selectin

prefers carbohydrate 6-sulfation totyrosine sulfation,

whereas p-selectin does not.

AUTHOR:

Kanamori Akiko; Kojima Naoya; Uchimura Kenji; Muramatsu Takashi; Tamatani Takuya; Berndt Michael C; Kansas Geoffrey

S; Kannaqi Reiji

CORPORATE SOURCE:

Program of Molecular Pathology, Aichi Cancer Center,

Research Institute, Nagoya 464-8681, Japan.

CONTRACT NUMBER:

HL55647 (NHLBI)

SOURCE:

JOURNAL OF BIOLOGICAL CHEMISTRY, (2002 Sep 6) 277 (36)

32578-86.

Journal code: 2985121R. ISSN: 0021-9258.

PUB. COUNTRY:

United States

DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

200210

ENTRY DATE:

Entered STN: 20020912

Last Updated on STN: 20030105

Entered Medline: 20021029

AB 1- and P-selectin are known to require sulfation in their ligand molecules. We investigated the significance of carbohydrate 6-sulfation and tyrosine sulfation in selectin-mediated cell adhesion. COS-7 cells were genetically engineered to express P-selectin glycoprotein ligand-1 (PSGL-1) or its mutant in various combinations with 6-O-sulfotransferase (6-Sul-T) and/or

alphal-->3fucosyltransferase VII (Fuc-T VII). The cells transfected with PSGL-1, 6-Sul-T, and Fuc-T VII cDNAs supported rolling mediated by all three selectins and provided the best experimental system so far to estimate kinetic parameters in selectin-mediated cell adhesion for all three selectins using the identical rolling substrate and to compare the ligand specificity of each selectin. L-selectin

-mediated rolling was drastically impaired if the cells lacked carbohydrate 6-sulfation elaborated by 6-Sul-T, but not affected when PSGL-1 was replaced with a mutant lacking three tyrosine residues at its NH(2) terminus. L-selectin-mediated adhesion was also hardly affected by meantagin treatment of the cells, which closved a

hardly affected by mocarhagin treatment of the cells, which cleaved a short peptide containing sulfated tyrosine residues from PSGL-1. In contrast, **P-selectin**-mediated rolling was abolished

when PSGL-1 was either mutated or cleaved by mocarhagin at its NH(2) terminus, whereas the cells **expressing** PSGL-1 and Fuc-T VII but not 6-Sul-T showed only a modest decrease in **P-selectin** 

-mediated adhesion. These results indicate that  $\mathbf{L}$ -selectin prefers carbohydrate 6-sulfation much more than tyrosine sulfation, whereas  $\mathbf{P}$ -selectin favors tyrosine

sulfation in the PSGL-1 molecule far more than carbohydrate 6-sulfation. E-selectin-mediated adhesion was sulfation-independent requiring only Fuc-T VII, and thus the three members of the selectin family have distinct requirements for ligand sulfation.

L19 ANSWER 3 OF 16 BIOTECHDS COPYRIGHT 2004 THOMSON DERWENT/ISI on STN ACCESSION NUMBER: 2002-06107 BIOTECHDS

TITLE:

New enzyme, useful for modifying acceptor molecule, comprises an isolated  ${f L-selectin}$ 

sulfotransferase-2 that directs expression
of L-selectin ligand antigen, MECA-79 in

Chinese hamster ovary cells, or intestinal GlcNAc 6-

sulfotransferase;

plasmid pcDNA1.1/LSST-2-mediated enzyme gene transfer and expression in host cell for recombinant protein production in CHO cell for Crohn disease, ulcerative colitis, skin inflammatory disorder, allergic contact dermatitis, psoriasis, Lichen planus, lymphoma, chronic pneumonia, delayed-type hypersensitivity reaction,

diabetes and hyperplastic thymus therapy

AUTHOR: FUKUDA M; YEH J; HIRAOKA N

PATENT ASSIGNEE: BURNHAM INST

PATENT INFO: WO 2001085177 15 Nov 2001 APPLICATION INFO: WO 2000-US15452 11 May 2000 PRIORITY INFO: US 2000-569320 11 May 2000

DOCUMENT TYPE: Patent LANGUAGE: English

OTHER SOURCE: WPI: 2002-075226 [10]

AB DERWENT ABSTRACT:

NOVELTY - An isolated polypeptide (I) comprising an amino acid sequence encoding L-selectin sulfotransferase-2

(LSST-2) or its active fragment that directs **expression** of a **L-selectin** ligand antigen, MECA-79 in Chinese hamster ovary (CHO) cells, or comprising a sequence encoding intestinal GlcNAc 6-sulfotransferase (I-GLcNAc6ST), is new.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following: (1) treating or preventing (M) an **L-selectin**-mediated condition in a subject, comprising reducing the **expression** or activity of a betal, 3-N-acetylglucosaminyl transferase (betal, 3GnT) that directs **expression** of a MECA-79 antigen; (2) an isolated **L-selectin** antagonist (II) comprising an extended core 1 structure comprising the oligosaccharide Galbetal-4(SO3-6)GlcNAcbetal-3Galbetal-3GalNAc; and (3) an isolated nucleic acid molecule (III) comprising a sequence encoding (I).

WIDER DISCLOSURE - Disclosed as new are the following: (A) an isolated polypeptide which contains an amino acid sequence encoding a betal, 3GnT, or its active fragment, that directs expression of a MECA-79 antigen in CHO cells; (B) a substantially purified antibody material that specifically binds LSST-2, I-GLcNAc6ST or betal, 3GnT; (C) an isolated antisense nucleic acid molecule which contains a sequence that specifically binds to a sequence comprising 1333 or 1937 base pairs fully defined in the specification; (D) an oligonucleotide which contains a nucleotide sequence having at least 10 contiguous nucleotides of a sequence comprising 1333 or 1937 base pairs fully defined in the specification, or a nucleotide sequence complementary to it; (E) a vector containing a nucleic acid molecule encoding LSST-2; and (F) a host cell containing the above-mentioned vector.

BIOTECHNOLOGY - Preferred Polypeptide: LSST-2 produces MECA-79 antigen, when co-transfected into CHO cells together with beta1, 3GnT. I-GLcNAc6ST in combination with beta1, 3GnT produces MECA-79 antigen in Lec-2 cells, but not in CHO cells. Preferred Antagonist: (II) comprises two or more of the oligosaccharide Galbeta1-4(SO3-6)GlcNAcbeta1-3Galbeta1-3GalNAc, or two or more of the oligosaccharide NeuNAcalpha2-3Galbeta1-4(sulfo-6(Fucalpha1-3)GlcNAc)beta1-3Galbeta1-3GalNAcalpha1. Preferred Method: (M) involves administering to the subject an oligosaccharide L-selectin antagonist that inhibits the binding of L-selectin to a MECA-79 antigen, an inhibitory antibody material that specifically binds beta1, 3GnT, or a beta1, 3GnT antisense nucleic acid molecule comprising 20 nucleotides complementary to a sequence of 1208 or 1337 base pairs fully defined in the specification. (M) further comprises reducing the expression or activity of LSST-2 in the subject.

ACTIVITY - Antiinflammatory; antiulcer; antipsoriatic; antidiabetic; dermatological; antiallergic. No biological data provided.

MECHANISM OF ACTION - Inhibitor of binding of L-selectin to MECA-79 antigen (claimed). Inhibition of L-selectin ligand antigen (MECA-79) antibody binding by synthetic oligosaccharide such as Galbeta1-4(SO3-6)GlcNAcbeta1-3Galbeta1-3GalNAc was tested. Synthetic oligosaccharides were mixed at the indicated concentrations with MECA-79 antibody. The mixtures were incubated at room temperature for one hour before addition to wells precoated with transfected media from CHO/CD34/FT7/LSST/core 1 extension beta1, 3-N-acetylglucosaminyl transferase (beta1, 3GnT) cells. Antibody

binding was assayed. The results showed that the 6-S-extended core 1 structure, Galbeta1-4(SO3-6)GlcNAcbeta1-3Galbeta1-3GalNAc, was active in inhibiting binding of anti-MECA-79 antibody.

USE - (M) is useful for treating or preventing an L-selectin-mediated condition in a subject (claimed). (I) is useful for modifying an acceptor molecule by contacting the acceptor molecule with (I) or its active fragment. (III) is useful for treating L-selectin mediated conditions such as Crohn's disease and ulcerative colitis, inflammatory disorders of the skin such as allergic contact dermatitis, psoriasis and Lichen planus, lymphomas, chronic pneumonia, delayed-type hypersensitivity reactions, diabetes and hyperplastic thymus.

hyperplastic thymus. ADMINISTRATION - No administration details are given. EXAMPLE - A nucleic acid molecule encoding human L -selectin ligand sulfotransferase-2 (LSST-2), which, together with the beta1-3-N-acetylglucosaminyl transferase (beta1, 3GnT), directed expression of the L-selectin ligand antigen, MECA-79, was isolated. Human genomic DNA was used as the template for polymerase chain reaction (PCR)-based cloning. Primers corresponding to nucleotides 891-910 and nucleotides 1327-1302 of mouse LSST-1 were used to amplify human genomic DNA. The amplified products were cloned into pBluescript by TA cloning. The resultant coding sequence was 79.2% identical to mouse LSST-1 at the nucleotide level. To clone the full-length LSST-2 coding sequence, a P1 phage library of a human genomic DNA was amplified using primers 5'-CCGAATTCTCCCGAGAACGCACAAG-3' and 5'-CCCAAGCTTCTCATAGAGCACAAGCAG-3. From the single positive clone, DNA was purified and sequenced directly. The coding sequence present on the single exon was confirmed by reverse transcriptase (RT)-PCR using poly(A)+ RNA isolated from human lymph node. Three pairs of primers used in these PCR reactions corresponded to 5'-TTGGCCAGAAGGGGAATAG-3' (S1), 5'-CCACTGAAAGAGGCTGGACTGT-3'(S2), 5'-GGTTCTGTCTTCCTGGCGCTC-3' (S3), 5'-TTTGGCAGATGACCTGCATCAC-3' (S4), 5'-AGAACGCACAAAGGAGATCTCA-3' (S5), and 5'-AGATGTAGGCAAGGCTCAGAAG-3'(S6). PCR with the S1 and S2 resulted in the expected characteristic fragment of 470 base pairs, PCR with S3 and S4 resulted in the expected characteristic fragment of 617 base pairs, and PCR with S5 and S6 resulted in the expected characteristic fragment of 600 base pairs. The cDNA containing full-length coding sequence of human LSST-2 was excised by XbaI and TfiI, blunt-ended and cloned into pcDNA1.1. The resulting LSST-2 expression vector, in which the LSST-2 coding sequence was expressed under control of the CMV promoter, was designated pcDNA1.1/LSST-2. (98 pages)

L19 ANSWER 4 OF 16 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER:

2001:64196 HCAPLUS

DOCUMENT NUMBER:

134:127828

TITLE:

Cloning of nucleic acid sequences encoding

human and murine glycosyl

sulfotransferases

INVENTOR(S):

Rosen, Steven D.; Lee, Jin Kyu; Hemmerich, Stefan

Regents of the University of California, USA

PATENT ASSIGNEE(S): SOURCE:

PCT Int. Appl., 128 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE
WO 2001006015 A1 20010125 WO 2000-US19741 20000719

W: AU, CA, JP

RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,

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EP 1210455
                       A1
                            20020605
                                           EP 2000-948806
                                                            20000719
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
                                           JP 2001-511223
                                                            20000719
     JP 2003505039
                            20030212
                                        US 1999-144694P P
PRIORITY APPLN. INFO.:
                                                            19990720
                                        US 2000-593828
                                                         A 20000713
                                        WO 2000-US19741 W 20000719
    Novel glycosyl sulfotransferases (GST-4.alpha., GST-4.beta., and
AΒ
     GST-6 from human; GST-4 and GST-6 from mouse) and polypeptides
     related thereto, as well as nucleic acid compns. encoding the same, are
     provided. The glycosyl sulfotransferases are type 2 membrane
     proteins having a relatively short transmembrane domain and N-terminal
     cytoplasmic tail of varying length, and are capable of sulfating selectin
     ligands, particularly L-selectin ligands (e.g.,
     GlyCAM-1). Genomic DNA sequences encoding human
     GST-4 and GST-6 and for mouse GST-6 are also provided. The subject
     polypeptides and nucleic acid compns. find use in a variety of
     applications, including various diagnostic and therapeutic agent screening
     applications. Also provided are methods of inhibiting selectin-mediated
     binding events and methods of treating disease conditions assocd.
     therewith, particularly by administering an inhibitor of at least one of
     GST-4.alpha., GST-4.beta., and GST-6.
REFERENCE COUNT:
                               THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS
                         2
                               RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
L19 ANSWER 5 OF 16 EMBASE COPYRIGHT 2004 ELSEVIER INC. ALL RIGHTS RESERVED.
     on STN
ACCESSION NUMBER:
                    2001194520 EMBASE
TITLE:
                    Structural and functional features of the CD34
                    antigen: An update.
AUTHOR:
                    Lanza F.; Healy L.; Sutherland D.R.
CORPORATE SOURCE:
                    D.R. Sutherland, University Health Network, Princess
                    Margaret Hospital, Dept. of Med. Oncology/Hematology, 610
                    University Avenue, Toronto, Ont. M5G 2M9, Canada.
                    rob.sutherland@utoronto.ca
SOURCE:
                    Journal of Biological Regulators and Homeostatic Agents,
                    (2001) 15/1 (1-13).
                    Refs: 95
                    ISSN: 0393-974X CODEN: JBRAER
COUNTRY:
                    Italy
                    Journal; General Review
DOCUMENT TYPE:
FILE SEGMENT:
                    025
                            Hematology
                    026
                            Immunology, Serology and Transplantation
                    029
                            Clinical Biochemistry
LANGUAGE:
                    English
SUMMARY LANGUAGE:
                    English
     CD34 is a heavily glycosylated type I transmembrane molecule,
     that can be phoshorylated by a variety of kinases including Protein kinase
     C and Tyrosine kinases. The classification of epitopes detected by
     different CD34 MAbs has aided the selection of appropriate
     antibodies for use in specific clinical and research laboratory settings.
     Detailed structural analyses and cloning studies have confirmed
     that CD34 is a sialomucin, and have suggested that the fine
     composition of the carbohydrate moieties contained in its extended
     N-terminal region is important in determining its interactions with a
     variety of different ligands. For high endothelial venules (HEV)
     CD34 to serve as a ligand for L-selectin, the
     O-linked glycans of HEV CD34 are modified in an exquisitely
     specific manner with a variety of sialyl- and sulfo-transferases. In
     contrast, CD34 is not the ligand for L-
     selectin in hematopoietic stem/progenitor cells (HSPCs) and
     despite much endeavour, ligands for hematopoietic CD34 remain to
```

PT, SE

be identified.

L19 ANSWER 6 OF 16 SCISEARCH COPYRIGHT 2004 THOMSON ISI on STN

ACCESSION NUMBER: 2001:39518 SCISEARCH

THE GENUINE ARTICLE: 386BD

TITLE: Sulfa

Sulfation of N-acetylglucosamine by chondroitin 6-

sulfotransferase 2 (GST-5)

AUTHOR: Bhakta S; Bartes A; Bowman K G; Kao W M; Polsky I; Lee J

K; Cook B N; Bruehl R E; Rosen S D; Bertozzi C R;

Hemmerich S (Reprint)

CORPORATE SOURCE: Thios Biotechnol, 828 Clayton St, San Francisco, CA 94117

USA (Reprint); Roche Biosci, Dept Resp Dis, Palo Alto, CA 94304 USA; Univ Calif Berkeley, Dept Chem, Berkeley, CA 94720 USA; Univ Calif Berkeley, Dept Mol & Cell Biol, Berkeley, CA 94720 USA; Univ Calif Berkeley, Howard Hughes

Med Inst, Berkeley, CA 94720 USA; Univ Calif San

Francisco, Dept Anat, San Francisco, CA 94143 USA; Univ Calif San Francisco, Program Immunol, San Francisco, CA

94143 USA

COUNTRY OF AUTHOR: USA

SOURCE: JOU

JOURNAL OF BIOLOGICAL CHEMISTRY, (22 DEC 2000) Vol. 275,

No. 51, pp. 40226-40234.

Publisher: AMER SOC BIOCHEMISTRY MOLECULAR BIOLOGY INC,

9650 ROCKVILLE PIKE, BETHESDA, MD 20814 USA.

ISSN: 0021-9258. Article; Journal

DOCUMENT TYPE: LANGUAGE:

English

REFERENCE COUNT:

43

\*ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS\*

AB Based on sequence homology with a previously cloned

human GlcNAc 6-O-sulfotransferase, we have identified an open reading frame (ORF) encoding a novel member of the Gal/GalNAc/GlcNAc 6-O-sulfotransferase (GST) family termed GST-5 on the

human X chromosome (band Xp11). GST-5 has recently been characterized as a novel GalNAc 6-0-sulfotransferase termed chondroitin 6-sulfotransferase-2 (Kitagawa, H., Fujita, M,,

chondroitin 6-sulfotransferase-2 (Kitagawa, H., Fujita, M,, Itio, N., and Sugahara K, (2000) J. Biol Chem. 275, 21075-21080), We have

coexpressed a human GST-5 cDNA with a GlyCAM-1

/IgG fusion protein in COS-7 cells and observed fourfold enhanced [S-35] sulfate incorporation into this mucin acceptor. All mucin-associated

[S-35] sulfate was incorporated as GlcNAc-6-sulfate or Gal

beta1-->4GlcNAc-6-sulfate. GST-5 was also **expressed** in soluble epitope-tagged form and found to catalyze 6-0-sulfation of GlcNAc residues in synthetic acceptor structures. In particular, GST-R was found to

in synthetic acceptor structures. In particular, GST-B was found to catalyze 6-O-sulfation of beta -benzyl GlcNAc but not alpha- or beta -benzyl GalNAc, In the mouse genome we have found a homologous ORF that

predicts a novel murine GlcNAc 6-O-sulfotransferase with 88% identity to the human enzyme. This gene was mapped to mouse

chromosome X at band XA3.1-3.2. GST-5 is the newest member of an emerging

family of carbohydrate 6-O-sulfotransferases that includes chondroitin g-sulfotransferase (GST-0), keratan-sulfate galactose 6-O-sulfotransferase (GST-1), the ubiquitously

expressed GlcNAc 6-0-sulfotransferase (GST-S), high
endothelial cell GlcNAc 6-0-sulfotransferase (GST-3), and

intestinal GlcNAc 6-O-sulfotransferase (GST-4),

L19 ANSWER 7 OF 16 SCISEARCH COPYRIGHT 2004 THOMSON ISI on STN

ACCESSION NUMBER: 2000:516591 SCISEARCH

THE GENUINE ARTICLE: 330AU

TITLE: Molecular cloning and expression of two distinct human chondroitin 4-0-

sulfotransferases that belong to the HNK-1

sulfotransferase gene family

AUTHOR: Hiraoka N; Nakagawa H; Ong E; Akama T O; Fukuda M N;

Fukuda M

BURNHAM INST, CTR CANC RES, GLYCOBIOL PROGRAM, 10901 N CORPORATE SOURCE:

TORREY PINES RD, LA JOLLA, CA 92037 (Reprint); BURNHAM INST, CTR CANC RES, GLYCOBIOL PROGRAM, LA JOLLA, CA 92037

COUNTRY OF AUTHOR:

USA

SOURCE:

JOURNAL OF BIOLOGICAL CHEMISTRY, (30 JUN 2000) Vol. 275,

No. 26, pp. 20188-20196.

Publisher: AMER SOC BIOCHEMISTRY MOLECULAR BIOLOGY INC,

9650 ROCKVILLE PIKE, BETHESDA, MD 20814.

ISSN: 0021-9258.

DOCUMENT TYPE:

Article; Journal

FILE SEGMENT:

LIFE

LANGUAGE:

English

63

REFERENCE COUNT:

\*ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS\*

Using an expression cloning strategy, the cDNA AB encoding the human HNK-1 sulfotransferase (HNK-1ST) has been cloned. During this cloning we found that HNK-1ST and other Golgi-associated sulfotransferases cloned before share homologous sequences including the RDP motif (Ong, E., Yeh, J.-C., Ding, Y., Hindsgaul, O., and Fukuda, M. (1998) J. Biol. Chem, 223, 5190-5195). Using this conserved sequence in HNK-1ST as a probe, we identified two  ${\bf expressed}$  sequence tags in  $\bar{{\tt EST}}$  data base which have 31.6 and 30.7% identity with HNK-1ST at the amino acid levels, Expression of these two full-length cDNAs failed to form HNK-1 glycan nor to add sulfate to CD34 or NCAM. Surprisingly, proteins expressed by these cDNAs transferred sulfate to the C-4 position of N-acetylgalactosamine in chondroitin and desulfated dermatan sulfate, thus we named these two enzymes, chondroitin 4-0-sulfotransferas 1 and -2 (C4ST-1 and C4ST-2). Both C4ST-1 and C4ST-2, however, did not

form 4,6-di-O-sulfated N-acetylgalactosamine when chondroitin sulfate C was used as an acceptor. Moreover, analysis of S-35-labeled dermatan  $\,$ sulfate formed by C4ST-1 indicate that sulfation preferentially took place in GlcA-->GalNAc unit than in IdoA-->GalNAc unit, suggesting that 4-O-sulfation at N-acetylgalactosamine may precede epimerization of glucuronic acid to iduronic acid during dermatan sulfate biosynthesis, Northern analysis demonstrated that the transcript for C4ST-1 is predominantly expressed in peripheral leukocytes and hematopoietic tissues while the C4ST-2 transcript is more widely

expressed in various tissues. These results indicate C4ST-1 and C4ST-2 play complementary roles in chondroitin and dermatan sulfate synthesis in different tissues.

L19 ANSWER 8 OF 16 SCISEARCH COPYRIGHT 2004 THOMSON ISI on STN

THE GENUINE ARTICLE: 356FA

2000:728342 SCISEARCH

TITLE:

Differential carbohydrate recognition of two GlcNAc-6-

sulfotransferases with possible roles in L

-selectin ligand biosynthesis

AUTHOR:

Cook B N; Bhakta S; Biegel T; Bowman K G; Armstrong J I;

CORPORATE SOURCE:

ACCESSION NUMBER:

Hemmerich S; Bertozzi C R (Reprint)

UNIV CALIF BERKELEY, DEPT CHEM, BERKELEY, CA 94720 (Reprint); UNIV CALIF BERKELEY, DEPT CHEM, BERKELEY, CA

94720; UNIV CALIF BERKELEY, DEPT MOL & CELL BIOL, BERKELEY, CA 94720; ROCHE BIOSCI, DEPT MOL BIOL, PALO

ALTO, CA 94304

COUNTRY OF AUTHOR:

USA

SOURCE:

JOURNAL OF THE AMERICAN CHEMICAL SOCIETY, (13 SEP 2000)

Vol. 122, No. 36, pp. 8612-8622.

Publisher: AMER CHEMICAL SOC, 1155 16TH ST, NW,

WASHINGTON, DC 20036.

ISSN: 0002-7863.

DOCUMENT TYPE:

Article; Journal

FILE SEGMENT:

PHYS; LIFE

LANGUAGE:

English

REFERENCE COUNT:

\*ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS\*

Two human GlcNAc-6-sulfotransferases, CHST2 and HEC-GlcNAc6ST, have been recently identified as possible contributors to the inflammatory response by virtue of their participation in Lselectin liquid biosynthesis. Selective inhibitors would facilitate their functional elucidation and might provide leads for antiinflammatory therapy. Here we investigate the critical elements of a disaccharide substrate that are required for recognition by CHST2 and HEC-GlcNAc6ST. A panel of disaccharide analogues, bearing modifications to the pyranose rings and aglycon substituents, were synthesized and screened for substrate activity with each enzyme. Both GlcNAc-6sulfotransferases required the 2-N-acetamido and 4-hydroxyl groups of a terminal GlcNAc residue for conversion to product. Both enzymes tolerated modifications to the reducing terminal pyranose. Key differences in recognition of an amide group in the aglycon substituent were observed, providing the basis for future glycomimetic inhibitor design.

L19 ANSWER 9 OF 16 EMBASE COPYRIGHT 2004 ELSEVIER INC. ALL RIGHTS RESERVED.

on STN

ACCESSION NUMBER:

2000362283 EMBASE

TITLE:

Sulfotransferases as targets for therapeutic

intervention.

AUTHOR:

Armstrong J.I.; Bertozzi C.R.

CORPORATE SOURCE:

C.R. Bertozzi, Department Chemistry, University of

California, Berkeley, CA 94720, United States.

bertozzi@cchem.berkeley.edu

SOURCE:

Current Opinion in Drug Discovery and Development, (2000)

3/5 (502-515).

Refs: 102

ISSN: 1367-6733 CODEN: CODDFF

COUNTRY:

United Kingdom

DOCUMENT TYPE: FILE SEGMENT:

Journal; General Review Pharmacology 030

037 Drug Literature Index

LANGUAGE:

English SUMMARY LANGUAGE: English

Sulfated biomolecules regulate a diverse array of normal and pathological cellular communication events. The participation of these bioconjugates in a variety of disease states has sparked interest in the enzyme class that installs the sulfate esters: the sulfotransferases. Recent advances in the cloning and characterization of sulfotransferase enzymes and our understanding of the role of sulfated biomolecules in disease states have prompted the search for specific sulfotransferase inhibitors. Evidence for the participation of sulfated carbohydrates and proteins in acute and chronic inflammation, tumor progression and microbial pathogenesis is presented herein, followed by a discussion of sulfotransferase mechanism

L19 ANSWER 10 OF 16 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER:

1999:626310 HCAPLUS

and approaches to inhibiting sulfotransferase activity.

DOCUMENT NUMBER:

131:254317

TITLE:

Cloning of human and murine

glycosylsulfotransferase-3 and its role in

selectin-mediated binding events

INVENTOR(S):

Bistrup, Annette; Rosen, Steven D.; Tangemann,

Kirsten; Hemmerich, Stefan

PATENT ASSIGNEE(S):

The Regents of the University of California, USA;

Syntex, Incorporated

SOURCE:

PCT Int. Appl., 60 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

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PATENT NO.
                      KIND DATE
                                           APPLICATION NO.
                                                            DATE
     WO 9949018
                            19990930
                                           WO 1999-US4316
                                                            19990226
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             KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN,
             MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM,
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             TJ, TM
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             CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
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                       Α1
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                                          AU 1999-27945
                                                             19990226
    AU 764852
                       B2
                            20030904
     EP 1062326
                       A1
                            20001227
                                           EP 1999-908538
                                                            19990226
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             IE, FI
     JP 2002507409
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                                           JP 2000-537979
                                                             19990226
PRIORITY APPLN. INFO.:
                                        US 1998-45284
                                                         Α
                                                            19980320
                                        US 1998-190911
                                                         A 19981112
                                        WO 1999-US4316
                                                         W 19990226
     Novel mammalian glycosylsulfotransferases expressed in high
AB
     endothelial cells (GST-3) and polypeptides related thereto, as well as
     nucleic acid compns. encoding the same, are provided. The novel mammalian
     enzyme is a type 2 membrane protein having a relatively short
     transmembrane domain and a short N-terminal cytoplasmic tail.
                                                                    GST-3 is
     capable of sulfating selectin ligands, particularly L-
     selectin ligands., e.g., GlyCam-1, and is
     predicted to have N-acetylglucosamine-6-O-sulfotransferase
     activity. Human GST-3 is 386 amino acids in length, is highly
     glycosylated, and its expression is highly restricted; for
     example, human GST-3 is expressed in high endothelial
     cells (HEC) but not tonsillar lymphocytes or primary cultured
    human umbilical vein endothelial cells. Mouse Gst-3 is a 388
     amino acid protein. Also provided are keratin sulfate galactosyl-6-
     sulfotransferase (KSGal6ST) homologs that are selectively
     expressed in HEC. The subject polypeptides and nucleic acid
     compns. find use in a variety of applications, including research,
     diagnostic, and therapeutic agent screening applications. Also provided
     are methods of inhibiting selectin-mediated binding events and methods of
     treating disease conditions assocd. therewith, particularly by
     administering an inhibitor of at least one of GST-3 or KSGal6ST, or
     homologs thereof.
REFERENCE COUNT:
                               THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS
                         1
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L19 ANSWER 11 OF 16
                         MEDLINE on STN
                    1999439774
ACCESSION NUMBER:
                                   MEDLINE
DOCUMENT NUMBER:
                               PubMed ID: 10510083
                    99439774
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TITLE: Sulfation of a high endothelial venule-expressed

ligand for L-selectin. Effects on tethering and rolling of lymphocytes.

AUTHOR: Tangemann K; Bistrup A; Hemmerich S; Rosen S D CORPORATE SOURCE: Department of Anatomy, Program in Immunology, and

Cardiovascular Research Institute, University of California

San Francisco, San Francisco, California 94143, USA.

CONTRACT NUMBER: R37GM23547 (NIGMS)

RO1GM5741 (NIGMS)

SOURCE: JOURNAL OF EXPERIMENTAL MEDICINE, (1999 Oct 4) 190 (7)

935-42.

Journal code: 2985109R. ISSN: 0022-1007.

PUB. COUNTRY:

United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

199911

ENTRY DATE:

Entered STN: 20000111

Last Updated on STN: 20000111 Entered Medline: 19991104

AB During lymphocyte homing, L-selectin mediates the

tethering and rolling of lymphocytes on high endothelial venules (HEVs) in

secondary lymphoid organs. The L-selectin ligands on

HEV are a set of mucin-like glycoproteins, for which glycosylation-

dependent cell adhesion molecule 1 (GlyCAM-1) is a

candidate. Optimal binding in equilibrium measurements requires sulfation, sialylation, and fucosylation of ligands. Analysis of

GlyCAM-1 has revealed two sulfation modifications

(galactose [Gal]-6-sulfate and N-acetylglucosamine [GlcNAc]-6-sulfate) of

sialyl Lewis x. Recently, three related sulfotransferases

(keratan sulfate galactose-6-sulfotransferase [KSGal6ST], high

endothelial cell N-acetylglucosamine-6-sulfotransferase

[GlcNAc6ST], and human GlcNAc6ST) were cloned, which

can generate Gal-6-sulfate and GlcNAc-6-sulfate in GlyCAM-

1. Imparting these modifications to GlyCAM-1,

together with appropriate fucosylation, yields enhanced rolling ligands for both peripheral blood lymphocytes and Jurkat cells in flow chamber assays as compared with those generated with exogenous fucosyltransferase. Either sulfation modification results in an increased number of tethered and rolling lymphocytes, a reduction in overall rolling velocity associated with more frequent pausing of the cells, and an enhanced resistance of rolling cells to detachment by shear. All of these effects are predicted to promote the overall efficiency of lymphocyte homing. In contrast, the rolling interactions of E-selectin transfectants with the same ligands are not affected by sulfation.

L19 ANSWER 12 OF 16 MEDLINE on STN

ACCESSION NUMBER:

1999264336 MEDLINE

DOCUMENT NUMBER:

99264336 PubMed ID: 10330415

TITLE: Sulfotransf

Sulfotransferases of two specificities function

in the reconstitution of high endothelial cell ligands for

L-selectin.

 ${\tt AUTHOR:}$ 

Bistrup A; Bhakta S; Lee J K; Belov Y Y; Gunn M D; Zuo F R;

Huang C C; Kannagi R; Rosen S D; Hemmerich S

CORPORATE SOURCE:

Department of Anatomy and Program in Immunology, University

of California, San Francisco, California 94143, USA.

CONTRACT NUMBER:

GM57411 (NIGMS)

R37 GM23547 (NIGMS) SOURCE: JOUR

GMS) JOURNAL OF CELL BIOLOGY, (1999 May 17) 145 (4) 899-910.

Journal code: 0375356. ISSN: 0021-9525.

PUB. COUNTRY:

United States

DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English

FILE SEGMENT:

Priority Journals

OTHER SOURCE:

GENBANK-AF131235; GENBANK-AF131236

ENTRY MONTH:

199907

ENTRY DATE:

Entered STN: 19990730

Last Updated on STN: 19990730 Entered Medline: 19990721

AB L-selectin, a lectin-like receptor, mediates rolling of lymphocytes on high endothelial venules (HEVs) in secondary lymphoid organs by interacting with HEV ligands. These ligands consist of a complex of sialomucins, candidates for which are glycosylation- dependent

cell adhesion molecule 1 (GlyCAM-1), CD34, and podocalyxin. The ligands must be sialylated, fucosylated, and sulfated for optimal recognition by L-selectin. Our previous structural characterization of GlyCAM-1 has demonstrated two sulfation modifications, Gal-6-sulfate and GlcNAc-6-sulfate in the context of sialyl Lewis x. We now report the cloning of a Gal-6-sulfotransferase and a GlcNAc-6sulfotransferase, which can modify GlyCAM-1 and CD34. The Gal-6-sulfotransferase shows a wide tissue distribution. In contrast, the GlcNAc-6-sulfotransferase is highly restricted to HEVs, as revealed by Northern analysis and in situ hybridization. Expression of either enzyme in Chinese hamster ovary cells, along with CD34 and fucosyltransferase VII, results in ligand activity, as detected by binding of an Lselectin/IgM chimera. When coexpressed, the two sulfotransferases synergize to produce strongly enhanced chimera binding.

L19 ANSWER 13 OF 16 BIOSIS COPYRIGHT 2004 BIOLOGICAL ABSTRACTS INC. on STN

ACCESSION NUMBER: 1999:212549 BIOSIS DOCUMENT NUMBER: PREV199900212549

TITLE: Culture characterization of differentiated high endothelial

venule cells from human tonsils.

Baekkevold, Espen S. [Reprint author]; Jahnsen, Frode L.; AUTHOR (S):

Johansen, Finn-Eirik; Bakke, Oddmund; Gaudernack, Gustav;

Brandtzaeg, Per; Haraldsen, Guttorm

CORPORATE SOURCE: LIIPAT, Rikshospitalet, N-0027, Oslo, Norway

SOURCE: Laboratory Investigation, (March, 1999) Vol. 79, No. 3, pp.

327-336. print.

CODEN: LAINAW. ISSN: 0023-6837.

DOCUMENT TYPE: Article LANGUAGE: English

ENTRY DATE: Entered STN: 26 May 1999

Last Updated on STN: 26 May 1999

High endothelial venules (HEV) are specialized vessels that support abundant lymphocyte emigration from peripheral blood into secondary lymphoid organs. HEV endothelial cells (HEVEC) exhibit particular structural and functional features, including secretion of the HEV-specific extracellular matrix protein hevin and an array of uniquely glycosylated counter-receptors for L-selectin expressed on lymphocytes. These ligands are collectively called the peripheral lymph node addressin (PNAd), originally defined by the monoclonal antibody MECA-79. PNAd expression was used to purify HEVEC by positive immunoselection from enzyme-digested human tonsils after negative immunoselection for other cells. Purified HEVEC maintained secretion of hevin and homogenous expression of intercellular adhesion molecule (ICAM)-1 (CD54), ICAM-2 (CD102), and CD31, at high levels following 8 days in culture. Expression of functional PNAd was maintained during the first 4 to 5 days of culture but decreased gradually and disappeared on day 8, while the expression of CD34 remained strong. However, the CD34 glycoform shifted toward the in situ phenotype of flat-walled vessels, suggesting that the observed loss of L-selectin binding determinants and MECA-79 antigen was due to down-regulation of the glycosyl- and sulfo-transferases essential for their expression. Our rapid and reproducible method to establish HEVEC cultures provides a useful mechanistic tool for identification of the factors that induce and maintain the HEV phenotype, as well as a source for isolation of HEV-specific genes.

L19 ANSWER 14 OF 16 BIOSIS COPYRIGHT 2004 BIOLOGICAL ABSTRACTS INC. on STN

ACCESSION NUMBER: 1999:175153 BIOSIS DOCUMENT NUMBER: PREV199900175153

TITLE: Cloning and characterization of two human carbohydrate sulfotransferases that are

expressed in high endothelial venules and confer

L-selectin binding activity onto recombinant L-selectin ligands.

AUTHOR (S): Bistrup, Annette; Tangemann, Kirsten; Bhakta, Sunil; Lee,

Jin Kyu; Belov, Yevgeniy Y.; Gunn, Michael Dee; Hemmerich,

Stefan; Rosen, Steven D.

Univ. California, San Francisco, CA 94143, USA CORPORATE SOURCE:

SOURCE: FASEB Journal, (March 12, 1999) Vol. 13, No. 4 PART 1, pp.

A313. print.

Meeting Info.: Annual Meeting of the Professional Research Scientists for Experimental Biology 99. Washington, D.C.,

USA. April 17-21, 1999.

CODEN: FAJOEC. ISSN: 0892-6638.

DOCUMENT TYPE: Conference; (Meeting)

Conference; Abstract; (Meeting Abstract)

LANGUAGE: English

ENTRY DATE: Entered STN: 5 May 1999

Last Updated on STN: 5 May 1999

L19 ANSWER 15 OF 16 MEDLINE on STN ACCESSION NUMBER: 1999361934 MEDLINE

DOCUMENT NUMBER: 99361934 PubMed ID: 10435581

A novel, high endothelial venule-specific TITLE: sulfotransferase expresses 6-sulfo sialyl

Lewis(x), an L-selectin ligand

displayed by CD34.

**AUTHOR:** Hiraoka N; Petryniak B; Nakayama J; Tsuboi S; Suzuki M; Yeh

J C; Izawa D; Tanaka T; Miyasaka M; Lowe J B; Fukuda M Glycobiology Program, Cancer Research Center, The Burnham CORPORATE SOURCE:

Institute, La Jolla, California 92037, USA.

CONTRACT NUMBER: PO1AI33189 (NIAID)

PO1CA71932 (NCI)

SOURCE: IMMUNITY, (1999 Jul) 11 (1) 79-89.

Journal code: 9432918. ISSN: 1074-7613.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals OTHER SOURCE: GENBANK-AF109155

ENTRY MONTH: 199908

ENTRY DATE: Entered STN: 19990820

Last Updated on STN: 19990820 Entered Medline: 19990811

AB L-selectin mediates lymphocyte homing by facilitating lymphocyte adhesion to unique carbohydrate liqands, sulfated sialyl Lewis(x), which are **expressed** on high endothelial venules (HEV) in secondary lymphoid organs. The nature of the sulfotransferase (s) that contribute to sulfation of such L-selectin counterreceptors has been uncertain. We herein describe a novel L -selectin ligand sulfotransferase, termed LSST, that directs the synthesis of the 6-sulfo sialyl Lewis(x) on Lselectin counterreceptors CD34, GlyCAM-

1, and MAdCAM-1. LSST is predominantly expressed in HEV and exhibits striking catalytic preference for core 2-branched mucin-type O-glycans as found in natural Lselectin counterreceptors. LSST enhances L-

selectin-mediated adhesion under shear compared to nonsulfated controls. LSST therefore corresponds to an HEV-specific sulfotransferase that contributes to the biosynthesis of L

-selectin ligands required for lymphocyte homing.

L19 ANSWER 16 OF 16 SCISEARCH COPYRIGHT 2004 THOMSON ISI on STN ACCESSION NUMBER: 1998:700521 SCISEARCH

THE GENUINE ARTICLE: 117KA

TITLE:

Human N-acetylglucosamine-6-0-

sulfotransferase involved in the biosynthesis of

6-sulfo sialyl Lewis X: Molecular cloning, chromosomal mapping, and expression in various

organs and tumor cells

**AUTHOR:** 

Uchimura K; Muramatsu H; Kaname T; Ogawa H; Yamakawa T; Fan Q W; Mitsuoka C; Kannaqi R; Habuchi O; Yokoyama I; Yamamura K; Ozaki T; Nakagawara A; Kadomatsu K; Muramatsu

T (Reprint)

CORPORATE SOURCE:

NAGOYA UNIV, SCH MED, DEPT BIOCHEM, SHOWA KU, 65 TSURUMAI CHO, NAGOYA, AICHI 4668550, JAPAN (Reprint); NAGOYA UNIV, SCH MED, DEPT BIOCHEM, SHOWA KU, NAGOYA, AICHI 4668550, JAPAN; NAGOYA UNIV, SCH MED, DEPT SURG 2, SHOWA KU, NAGOYA, AICHI 4668550, JAPAN; NAGOYA UNIV, SCH MED, DEPT INTERNAL MED 3, SHOWA KU, NAGOYA, AICHI 4668550, JAPAN; KUMAMOTO UNIV, SCH MED, INST MOL EMBRYOL & GENET, DEPT DEV GENET, KUMAMOTO 8620976, JAPAN; AICHI CANC CTR, RES INST, PROGRAM EXPT PATHOL, NAGOYA, AICHI 4640021, JAPAN; AICHI UNIV EDUC, DEPT LIFE SCI, KARIYA, AICHI 4488542, JAPAN; CHIBA CANC CTR, INST RES, DIV BIOCHEM, CHUOH KU, CHIBA

2600801, JAPAN

COUNTRY OF AUTHOR:

JAPAN

SOURCE:

JOURNAL OF BIOCHEMISTRY, (SEP 1998) Vol. 124, No. 3, pp.

670-678.

Publisher: JAPANESE BIOCHEMICAL SOC, ISHIKAWA BLDG-3F,

25-16 HONGO-5-CHOME, BUNKYO-KU, TOKYO 113, JAPAN.

ISSN: 0021-924X. Article; Journal

DOCUMENT TYPE: FILE SEGMENT:

LIFE

LANGUAGE:

English

REFERENCE COUNT:

system.

35

\*ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS\* AB N-Acetylglucosamine-6-0-sulfotransferase catalyzes the transfer of sulfate from 3'-phosphoadenosine 5'-phosphosulfate to position 6 of a non-reducing N-acetylglucosamine (GlcNAc) residue. We have cloned human GlcNAc-6-O-sulfotransferase cDNA, based on the sequence homology to cloned cDNA of mouse GlcNAc-6-0-sulfotransferase. The predicted protein sequence of the human enzyme was highly homologous to that of the mouse enzyme; in the 363 amino acid stretch of the catalytic region, the two proteins were nearly identical except for conservative changes in 3 amino acid residues. The expressed enzyme transferred sulfate to GlcNAc beta 1-3Gal beta 1-4GlcNAc beta 1-3Gal beta 1-4GlcNAc. Co-transfection of the enzyme cDNA and fucosyltransferase VII cDNA into COS-7 cells resulted in cell surface expression of 6-sulfo sialyl Lewis X. Fluorescence irt situ hybridization analysis revealed that the GlcNAc-6-O-sulfotransferase gene is located on human chromosome 7q31. mRNA of the human enzyme was strongly expressed in the bone marrow, peripheral blood leukocytes, spleen, brain, spinal cord, ovary, and placenta, and moderate levels of expression were observed in many organs including lymph nodes and thymus. In situ hybridization with the mouse system showed that the transcript was localized in specific regions of the brain, i.e. pyramidal cells in the CA3 subregion of the hippocampus, cerebellar nucleus and Purkinje cells. Among human tumor cells, strong expression of the mRNA was found in MOLT-4 and Jarkat lymphoblastic leukemia cells, Raji lymphoma cells, K-562 chronic myelogeneous leukemia cells, U251 glioma cells, and G361 melanoma cells. Carbohydrate structures synthesized by the sulfotransferase may

be involved in various aspects of the differentiation and behavior of blood cells, their progenitor cells, and neurons in the central nervous (FILE 'HOME' ENTERED AT 11:10:43 ON 07 JAN 2004)

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FILE 'MEDLINE, EMBASE, BIOSIS, BIOTECHDS, SCISEARCH, HCAPLUS, NTIS,
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              8 DUP REM L1 (3 DUPLICATES REMOVED)
L2
L3
              2 S "GST4 ALPHA"
              1 DUP REM L3 (1 DUPLICATE REMOVED)
L4
L5
             53 S "GST4"
             10 S HUMAN (A) L5
L6
Ь7
              2 DUP REM L6 (8 DUPLICATES REMOVED)
Г8
          13789 S SULFOTRANSFERASE?
L9
           5827 S HUMAN AND L8
L10
        6311680 S CLON? OR EXPRESS? OR RECOMBINANT
L11
           2771 S L9 AND L10
L12
          13955 S "L-SELECTIN"
L13
          19920 S "P-SELECTIN"
L14
          31224 S L12 OR L13
L15
            124 S L11 AND L14
             77 DUP REM L15 (47 DUPLICATES REMOVED)
L16
          59143 S "GLYCAM-1" OR "CD34" OR "MADCAM-1" OR "SGP200"
L17
L18
             16 S L16 AND L17
             16 DUP REM L18 (0 DUPLICATES REMOVED)
L19
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                   ROSEN RT/AU
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E2
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E3
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E6
                   ROSEN S C/AU
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E11
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                   ROSEN S F/AU
E12
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          2356 "ROSEN S"/AU
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                   LEE J A C/AU
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E9
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E10
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E11
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=> s e3
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=> e hemmerich s/au
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E2
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77
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             1
E6
             1
                   HEMMERICK PETER/AU
E7
             8
                   HEMMERLE A/AU
E8
             9
                   HEMMERLE A V/AU
E9
             6
                   HEMMERLE ANKE/AU
E10
                   HEMMERLE C/AU
E11
                   HEMMERLE CHRISTINE/AU
E12
=> s e3
           118 "HEMMERICH S"/AU
L22
=> d his
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L2
              8 DUP REM L1 (3 DUPLICATES REMOVED)
L3
              2 S "GST4 ALPHA"
L4
              1 DUP REM L3 (1 DUPLICATE REMOVED)
L5
             53 S "GST4"
L6
             10 S HUMAN (A) L5
L7
              2 DUP REM L6 (8 DUPLICATES REMOVED)
L8
          13789 S SULFOTRANSFERASE?
L9
           5827 S HUMAN AND L8
L10
        6311680 S CLON? OR EXPRESS? OR RECOMBINANT
L11
           2771 S L9 AND L10
L12
          13955 S "L-SELECTIN"
L13
          19920 S "P-SELECTIN"
L14
          31224 S L12 OR L13
L15
            124 S L11 AND L14
L16
             77 DUP REM L15 (47 DUPLICATES REMOVED)
L17
          59143 S "GLYCAM-1" OR "CD34" OR "MADCAM-1" OR "SGP200"
L18
             16 S L16 AND L17
T.19
             16 DUP REM L18 (0 DUPLICATES REMOVED)
                E ROSEN S/AU
           2356 S E3
L20
                E LEE J/AU
          13300 S E3
L21
                E HEMMERICH S/AU
L22
            118 S E3
=> s 121 or 120 or 122
         15770 L21 OR L20 OR L22
L23
=> s.15 and 123
L24
             3 L5 AND L23
=> dup rem 124
PROCESSING COMPLETED FOR L24
L25
              1 DUP REM L24 (2 DUPLICATES REMOVED)
=> d all
     ANSWER 1 OF 1
                       MEDLINE on STN
                                                          DUPLICATE 1
AN
     2001205848
                    MEDLINE
DN
              PubMed ID: 11181564
     Chromosomal localization and genomic organization for the galactose/
     N-acetylgalactosamine/N-acetylqlucosamine 6-0-sulfotransferase gene
     family.
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118 --> HEMMERICH S/AU

E3

ΑU Hemmerich S; Lee J K; Bhakta S; Bistrup A; Ruddle N R; Rosen S D CS Department of Respiratory Diseases, Roche Bioscience, Palo Alto, CA 94304, NC RO1GM5741 (NIGMS) SO GLYCOBIOLOGY, (2001 Jan) 11 (1) 75-87. Journal code: 9104124. ISSN: 0959-6658. CY England: United Kingdom DTJournal; Article; (JOURNAL ARTICLE) LA English FS Priority Journals OS GENBANK-AF176838; GENBANK-AF280086; GENBANK-AF280087; GENBANK-AF280088; GENBANK-AF280089; GENBANK-AI824100 EM 200106 ED Entered STN: 20010611 Last Updated on STN: 20010611 Entered Medline: 20010607 AB The galactose/N-acetylgalactosamine/N-acetylglucosamine 6-O-sulfotransferases (GSTs) are a family of Golgi-resident enzymes that transfer sulfate from 3'phosphoadenosine 5'phospho-sulfate to the 6-hydroxyl group of galactose, N-acetylgalactosamine, or N-acetylglucosamine in nascent glycoproteins. These sulfation modifications are functionally important in settings as diverse as cartilage structure and lymphocyte homing. To date six members of this gene family have been described in human and in mouse. We have determined the chromosomal localization of these genes as well as their genomic organization. While the broadly expressed enzymes implicated in proteoglycan biosynthesis are located on different chromosomes, the highly tissue specific enzymes GST-3 and 4 are encoded by genes located both in band q23.1--23.2 on chromosome 16. In the mouse, both genes reside in the syntenic region 8E1 on chromosome 8. This cross-species conserved clustering is suggestive of related functional roles for these genes. human GST4 locus actually contains two highly similar open reading frames (ORF) that are 50 kb apart and encode two highly similar enzyme isoforms termed GST-4 alpha and GST-4 beta. All genes except GST0 (chondroitin 6-O-sulfotransferase) contain intron-less ORFs. With one exception these are fused directly to sequences encoding the 3' untranslated regions (UTR) of the respective mature mRNAs. The 5' UTRs of these mRNAs are usually encoded by a number of short exons 5' of the respective ORF. 5'UTRs of the same enzyme expressed in different cell types are sometimes derived from different exons located upstream of the ORF. The genomic organization of the GSTs resembles that of certain glycosyltransferase gene families. Check Tags: Animal; Human; Support, Non-U.S. Gov't; Support, U.S. Gov't, P.H.S. Amino Acid Sequence Base Sequence Chromosome Mapping Chromosomes, Artificial, Bacterial \*Chromosomes, Human, Pair 16 Cloning, Molecular DNA, Complementary Glutathione Transferase: GE, genetics In Situ Hybridization, Fluorescence Mice

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(FILE 'HOME' ENTERED AT 11:10:43 ON 07 JAN 2004)

Molecular Sequence Data

2.5.1.18 (Glutathione Transferase)

FILE 'MEDLINE, EMBASE, BIOSIS, BIOTECHDS, SCISEARCH, HCAPLUS, NTIS,

0 (Chromosomes, Artificial, Bacterial); 0 (DNA, Complementary); EC

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LIFESCI' ENTERED AT 11:11:16 ON 07 JAN 2004
L1
             11 S GLYCOSYL (A) SULFOTRANSFERASE?
L2
              8 DUP REM L1 (3 DUPLICATES REMOVED)
L3
              2 S "GST4 ALPHA"
             1 DUP REM L3 (1 DUPLICATE REMOVED)
L5
             53 S "GST4"
             10 S HUMAN (A) L5
             2 DUP REM L6 (8 DUPLICATES REMOVED)
L7
          13789 S SULFOTRANSFERASE?
L8
L9
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        6311680 S CLON? OR EXPRESS? OR RECOMBINANT
L11
          2771 S L9 AND L10
          13955 S "L-SELECTIN"
L12
          19920 S "P-SELECTIN"
L13
L14
          31224 S L12 OR L13
L15
            124 S L11 AND L14
L16
             77 DUP REM L15 (47 DUPLICATES REMOVED)
L17
          59143 S "GLYCAM-1" OR "CD34" OR "MADCAM-1" OR "SGP200"
L18
             16 S L16 AND L17
L19
             16 DUP REM L18 (0 DUPLICATES REMOVED)
                E ROSEN S/AU
L20
           2356 S E3
                E LEE J/AU
          13300 S E3
L21
                E HEMMERICH S/AU
            118 S E3
L22
L23
          15770 S L21 OR L20 OR L22
L24
              3 S L5 AND L23
L25
              1 DUP REM L24 (2 DUPLICATES REMOVED)
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	Issue Date	Pages	Document ID	Title
1	20030501	78	US 20030082511 A1	Identification of modulatory molecules using inducible promoters
2	20020214	22	US 20020019019 A1	Method and apparatus for assaying a drug candidate to estimate a pharmacokinetic parameter associated therewith

	Issue Date	Pages	Document II	Title
1	20030911	34	US 2003017026 A1	Expression system
2	20030501	78	US 2003008251 A1	Identification of modulatory molecules using inducible promoters
3	20030213	33	US 2003003168 A1	Combined growth factor-deleted and thymidine kinase-deleted vaccinia virus vector
4	20020214	22	US 2002001901 A1	Method and apparatus for assaying a drug candidate to estimate a pharmacokinetic parameter associated therewith
5	20000711	14	US 6088277 A	Read only memory capable of realizing a high-speed read operation
6	19861216	22	US 4630188 A	Multi-zone ramp system for digital pulse generator and large scale integrated chip embodying the same
7	19821102	9	US 4357584 A	Acoustic wave devices

	Issue Date	Pages	Document ID	Title
1	20040101	106	US 20040002067 A1	Breast cancer progression signatures
2	20030814	278	US 20030154032 A1	Methods and compositions for diagnosing and treating rheumatoid arthritis
3	20030417	: 7 ク	US 20030073100 A1	Method of identifying renalgenerative agents using differential gene expression
4	20021107	:	US 20020164748 A1	Glycosyl sulfotransferase-3
5	20020604	17	US 6399358 B1	Human gene encoding human chondroitin 6-sulfotransferase
6	20020402	38	US 6365365 B1	Method of determining whether an agent modulates glycosyl sulfotransferase-3

2 20030306 202 US 20030044783 Human genes and gene expression products  3 20021107 36 US 20020164748 Glycosyl sulfotransferase-3  4 20011213 27 US 20010051370 Glycosyl sulfotransferase-3  5 20021105 179 US 6476195 B1 Secreted protein HNFGF20  5 20020402 38 US 6365365 B1 Method of determining whether an agent modulates glycosyl sulfotransferase-3		Issue Date	Pages	Document ID	Title
20021107 36 US 20020164748 Glycosyl sulfotransferase-3 4 20011213 27 US 20010051370 Glycosyl sulfotransferase-3 5 20021105 179 US 6476195 B1 Secreted protein HNFGF20 6 20020402 38 US 6365365 B1 Method of determining whether an agent modulates glycosyl sulfotransferase-3	1	20030731	104	;	Drug metabolizing enzymes
A1  Glycosyl sulfotransferase-3  US 20010051370 Glycosyl sulfotransferase-3  Secreted protein HNFGF20  Method of determining whether an agent modulates glycosyl sulfotransferase-3  Sulfotransferase-3  Method of determining whether an agent modulates glycosyl sulfotransferase-3	2	20030306	202		
A1  Glycosyl sulfotransferase-3  A1  Secreted protein HNFGF20  Method of determining whether an agent modulates glycosyl sulfotransferase-3  us 6365365 B1  Method of determining whether an agent modulates glycosyl sulfotransferase-3	3	20021107	36	•	Glycosyl sulfotransferase-3
Method of determining whether 20020402 38 US 6365365 B1 an agent modulates glycosyl sulfotransferase-3	4	20011213	27		Glycosyl sulfotransferase-3
6 20020402 38 US 6365365 B1 an agent modulates glycosyl sulfotransferase-3	5	20021105	179	US 6476195 B1	Secreted protein HNFGF20
7 20010724 27 US 6265192 B1 Glycosly sulfortransferase-3	6	20020402	38	US 6365365 B1	an agent modulates glycosyl
	7	20010724	27	US 6265192 B1	Glycosly sulfortransferase-3

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2	L3	2	l1 same l2
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6	L6	537	sulfotransferase\$2
7	L7	203	l1 same 16
8	L8	588758	clon\$3 or express\$3 or recombinant
9	L9	132	17 same 18
10	L10	3301 -	selectin
11	L11	6	19 same 110
12	L12	2665	rosen.in.

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	13	L13	55556	lee.in.
·	14	L14	68	hemmerich.in.
	15	L15	58256	l12 or l13 or l14
	16	L16	7	19 and 115